

characterization of samples of interest. For example, frequency dependent changes in either the electric or magnetic polarization of samples can be used to identify the type of sample, using knowledge of the behavior of various materials in electromagnetic fields from conventional solid state physics. These changes may be characterized over a broad range of frequencies. Accordingly, in one embodiment, by sweeping the RF frequency of signals applied to field-generating components (or using more sophisticated signal processing techniques), the frequency response (e.g., absorption spectrum) of the sample can be measured at a particular location, and the sample may be identified or characterized based on the measured response.

[0095] In yet other embodiments relating to the application of RF fields and sensing of field/sample interactions under the control of the RF/detection components 480, an RF field can be used to conduct local measurements of magnetic resonance in a uniform magnetic field applied to a sample. In particular, the spins or magnetic domains of a given sample oscillate with characteristic frequencies, which can be used to identify the type of spin or the sample itself. Magnetic resonance types include ferromagnetic resonance (FMR) (small YIG spheres can be used as magnetic beads, wherein a YIG sphere has a single magnetic domain that rotates freely at GHz frequencies because the bead is spherical). Additionally, Electron Spin Resonance (ESR) techniques may be employed to identify the g-factor of the spins involved to characterize their origin (i.e., the sample), as well as Nuclear Magnetic Resonance (NMR) to identify the g-factors of the nuclear spins. Thus, according to the principles discussed herein, a Magnetic Resonance Imaging (MRI) system may be implemented on a chip.

[0096] While not explicitly shown in FIGS. 1 and 2, according to various embodiments the field control components 400 also may include one or more analog to digital (A/D) and digital to analog (D/A) converters to facilitate the communication of various data and signals amongst other field control components, as well as to and from the IC chip 102. The field control components also may include digital signal processing components and signal amplification components to facilitate processing and transport of signals. Furthermore, the field control components may include a wireless transceiver and an antenna to facilitate wireless communication to and from the IC chip 102. In one exemplary wireless implementation, the ISM radio bands (free, non-commercial radio bands allowed for industrial, scientific and medical purposes) may be utilized for wireless communications between the IC chip 102 and a remote user or control interface (e.g., the one or more processors 600). Present wireless transceiver technology allows miniature, low-power transceivers to transmit and receive data at high data rates (e.g., several kilobits or megabits per second), which is sufficient for the reliable transfer of information to and from the IC chip 102.

[0097] Finally, FIGS. 1 and 2 also illustrate that the hybrid system 100 may include temperature regulation components 500 to facilitate biocompatibility of the hybrid system. For example, according to one embodiment, the temperature of the system may be regulated at or near a particular temperature to facilitate biocompatibility of the system with the samples under investigation. In one exemplary implementation, the temperature regulation components may include one or more “on-chip” temperature

sensors 500A (e.g., in proximity to the microfluidic system 300, as shown in FIG. 2) and an “off-chip” temperature controller 500B (e.g., a thermoelectric or “TE” cooler attached to the package substrate 110, as shown in FIG. 2). In one aspect, the one or more on-chip temperature sensors 500A sense the temperature of the IC chip in proximity to the microfluidic system and the one or more processors 600 compare the measured temperature to a reference temperature (e.g., 37° C.). The one or more processors in turn send an appropriate feedback control signal to the off-chip temperature controller 500B, which heats up or cools down the whole substrate accordingly. Temperature regulation components 500 are discussed further below in Section IV.

[0098] Having provided a general overview of a hybrid system according to the present disclosure for manipulation, detection, imaging and characterization of samples using electromagnetic fields, more detailed descriptions of various concepts related to different portions of the hybrid system, as well as some exemplary applications for such a system, are set forth below.

[0099] II. Microcoil Array

[0100] FIG. 6(a) is a conceptual perspective illustration of a microcoil array 200B that may be employed as field-generating components 200 in the hybrid system 100 shown in FIGS. 1 and 2, according to one embodiment of the present disclosure. In the example of FIG. 6(a), the array 200B includes five columns and five rows of essentially identical microcoils 212. Although FIG. 6(a) illustrates a five-by-five microcoil array, it should be appreciated that microcoil arrays according to various embodiments of the invention are not limited in this respect, and may have different numbers of microcoils and different geometric arrangements.

[0101] Like the microelectromagnet wire matrix 200A discussed above in connection with FIGS. 3(a)-(d), a microcoil array 200B similar to that shown in FIG. 6(a) may be configured and controlled to facilitate the manipulation of magnetic samples contained in the microfluidic system 300, including cells coupled to magnetic beads. FIG. 6(b) shows a conceptual illustration of a top (overhead) view of a portion of the array 200B shown in FIG. 6(a), looking down to the array through a portion of a microfluidic system 300 (e.g., a channel) that contains a liquid 306 in which are suspended exemplary samples 116 comprising a magnetic bead 112 attached to a cell 114 (i.e., a bead-bound cell). The liquid 306 also may contain one or more cells 114 that are not attached to a magnetic bead. In one embodiment, to manipulate the bead-bound cells 116 (or other types of magnetic samples), each microcoil 212 of the array 200B is independently connectable (via switching and multiplexing components, as discussed further below in connection with FIG. 13) to a source of controllable current. Thus, by independently controlling the magnitude of current flowing through each microcoil, various magnetic field patterns can be generated in proximity to the microcoil array 200B and employed to trap and otherwise manipulate magnetic samples.

[0102] As compared to the microelectromagnet wire matrix 200A, the microcoil array 200B generally is more efficient for at least some of the following exemplary reasons. First, the fields generated in the microcoil array are more highly localized than in the microelectromagnet wire